

Journal of the Royal Society of Medicine; 2015, Vol. 108(8) 297–298

To submit a letter in response to a JRSM article, please visit http://jrsm.sagepub.com to find the article concerned, then click on the 'Submit a comment' link found in the Reader Responses section. Comments will automatically be submitted for consideration to be published in print.

'Circumcision pain' unlikely to cause autism

Claims by Frisch and Simonsen¹ that 'circumcision pain' increases risk of autism spectrum disorder are flawed. They dredged Danish medical records of 342,877 boys aged 0–10 years looking for an adverse effect of circumcision. Marginal significance was apparent for autism spectrum disorder at age 0–4 years for 28 Muslim boys circumcised before the age of 2 years. Since HR was 1.54 (95% CI, 1.03–2.31), the circumcision effect involved approximately 10 boys. In 0–4-year-old circumcised non-Muslim boys they noted six autism spectrum disorder diagnoses and three hyperkinetic disorder diagnoses.

Of 337 Muslim boys aged 0–10 years with autism spectrum disorder, only 10.9% were 'circumcised', 89.1% being 'intact', which is improbable and undermines their findings.

The authors cite a survey that found pain was high in 4% of infants during six weeks post-circumcision. Since the Danish study was about pain why didn't Frisch and Simonsen examine other painful conditions? Urinary tract infections are associated with excruciating pain, are common and very much higher in uncircumcised male infants and boys.² If the authors' pain hypothesis were correct, then autism spectrum disorder should be associated with urinary tract infections and therefore being uncircumcised.

Anaesthetics are neurotoxic to the developing brain. Systemic use in children aged under 3 years is associated with later cognitive impairment.³ Could unnecessary general anaesthesia, not pain, have contributed to autism spectrum disorder?

They cite a study correlating autism spectrum disorder and circumcision prevalence post-1995, but fail to state that this was actually a study of paracetamol usage, circumcisions being merely a proxy.⁴ That 'hypothesis generating exploratory analysis' was prompted by the difference in paracetamol metabolism in immature brains that might produce neuronal damage in susceptible infants.

Most likely, both autism spectrum disorder diagnosis and early circumcision reflect parental conscientiousness.

Declarations

Competing interests: None declared

References

- 1. Frisch M and Simonsen J. Ritual circumcision and risk of autism spectrum disorder in 0- to 9-year old boys: national cohort study in Denmark. *J R Soc Med* 2015; Epub ahead of print Jan 8, 2015:doi:10.1177/0141076814565942.
- 2. Morris BJ and Wiswell TE. Circumcision and lifetime risk of urinary tract infections: A systematic review and meta-analysis. *J Urol* 2013; 189: 2118–2124.
- Ing C, DiMaggio C, Whitehouse A, Hegarty MK, Brady J, von Ungern-Sternberg BS, et al. Long-term differences in language and cognitive function after childhood exposure to anesthesia. *Pediatrics* 2012; 130: e476–e85.
- Bauer AZ and Kriebel D. Prenatal and perinatal analgesic exposure and autism: an ecological link. *Environ Health* 2013; 12: 41.

Brian J Morris

School of Medical Sciences, University of Sydney, Sydney, New South Wales, Australia Email: brian.morris@sydney.edu.au

Thomas E Wiswell

Center for Neonatal Care, Florida Hospital, Orlando, FL, USA

Circumcision-autism link needs thorough evaluation: Response to Morris and Wiswell

Morris and Wiswell express concern over small numbers of circumcised boys with autism spectrum disorder (ASD) and hyperkinetic disorder in non-Muslim families, apparently unaware of the fact that our findings were statistically significant despite the limited statistical power of these analyses, not because of it. Also, without substantiation, our critics consider the overall proportion of boys in Muslim families who were ritually circumcised by Danish doctors (10.9%) 'improbable', although no one knows the exact proportions of ritual circumcisions

performed by doctors vs. religious circumcisers in these boys. As explained, any mischaracterisation of boys as intact as opposed to circumcised would likely be non-differential, leading to conservative, not exaggerated, risk estimates.

We did not dredge medical records searching for adverse effects of circumcision. Rather, using national data, we tested the *a priori* hypothesis that ritual circumcision might be associated with an increased risk of ASD, based on converging evidence from animal, clinical and ecological studies. Morris and Wiswell's speculations about urinary tract infections and general anaesthesia in infants do not address our research question of whether circumcision is linked to an increased risk of ASD; and the paracetamol hypothesis was discussed in our article.

Morris and Wiswell claim that our study was 'flawed', but they do not provide relevant reasons for concern that were not already addressed in our original study.

Declarations

Competing interests: MF has been an author of articles on health-related and sexual consequences of male circumcision and has taken part in national and international debates on the ethics of male and female circumcision.

Reference

 Frisch M and Simonsen J. Ritual circumcision and risk of autism spectrum disorder in 0- to 9-year-old boys: national cohort study in Denmark. J R Soc Med 2015; DOI: 10.1177/0141076814565942.

Morten Frisch

Department of Epidemiology Research, Division of National Health Surveillance and Research, Statens Serum Institut, DK-2300 Copenhagen S, Denmark and Department of Clinical Medicine, Center for Sexology Research, Aalborg University, DK-9000 Aalborg, Denmark

Email: MFR@ssi.dk

Jacob Simonsen

Department of Epidemiology Research, Division of National Health Surveillance and Research, Statens Serum Institut, DK-2300 Copenhagen S, Denmark

